UNITED STATES DEPARTMENT OF COMMERCE United States Patent and Trademark Office Address: COMMISSIONER FOR PATENTS P.O. Box 1450 Alexandria, Virginia 22313-1450 www.uspto.gov

## NOTICE OF ALLOWANCE AND FEE(S) DUE

23389

7590

12/31/2008

SCULLY SCOTT MURPHY & PRESSER, PC 400 GARDEN CITY PLAZA SUITE 300 GARDEN CITY, NY 11530 EXAMINER

HOPKINS, CHRISTINE D

ART UNIT PAPER NUMBER

3735

DATE MAILED: 12/31/2008

APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
10/705,614	11/10/2003	James F. Hainfeld	16049Z	4692

TITLE OF INVENTION: METHODS OF ENHANCING RADIATION EFFECTS WITH METAL NANOPARTICLES

APPLN. TYPE	SMALL ENTITY	ISSUE FEE DUE	PUBLICATION FEE DUE	PREV. PAID ISSUE FEE	TOTAL FEE(S) DUE	DATE DUE
nonprovisional	YES	\$755	\$300	\$0	\$1055	03/31/2009

THE APPLICATION IDENTIFIED ABOVE HAS BEEN EXAMINED AND IS ALLOWED FOR ISSUANCE AS A PATENT. PROSECUTION ON THE MERITS IS CLOSED. THIS NOTICE OF ALLOWANCE IS NOT A GRANT OF PATENT RIGHTS. THIS APPLICATION IS SUBJECT TO WITHDRAWAL FROM ISSUE AT THE INITIATIVE OF THE OFFICE OR UPON PETITION BY THE APPLICANT. SEE 37 CFR 1.313 AND MPEP 1308.

THE ISSUE FEE AND PUBLICATION FEE (IF REQUIRED) MUST BE PAID WITHIN THREE MONTHS FROM THE MAILING DATE OF THIS NOTICE OR THIS APPLICATION SHALL BE REGARDED AS ABANDONED. THIS STATUTORY PERIOD CANNOT BE EXTENDED. SEE 35 U.S.C. 151. THE ISSUE FEE DUE INDICATED ABOVE DOES NOT REFLECT A CREDIT FOR ANY PREVIOUSLY PAID ISSUE FEE IN THIS APPLICATION. IF AN ISSUE FEE HAS PREVIOUSLY BEEN PAID IN THIS APPLICATION (AS SHOWN ABOVE), THE RETURN OF PART B OF THIS FORM WILL BE CONSIDERED A REQUEST TO REAPPLY THE PREVIOUSLY PAID ISSUE FEE TOWARD THE ISSUE FEE NOW DUE.

#### HOW TO REPLY TO THIS NOTICE:

I. Review the SMALL ENTITY status shown above.

If the SMALL ENTITY is shown as YES, verify your current SMALL ENTITY status:

A. If the status is the same, pay the TOTAL FEE(S) DUE shown above.

B. If the status above is to be removed, check box 5b on Part B - Fee(s) Transmittal and pay the PUBLICATION FEE (if required) and twice the amount of the ISSUE FEE shown above, or

If the SMALL ENTITY is shown as NO:

A. Pay TOTAL FEE(S) DUE shown above, or

B. If applicant claimed SMALL ENTITY status before, or is now claiming SMALL ENTITY status, check box 5a on Part B - Fee(s) Transmittal and pay the PUBLICATION FEE (if required) and 1/2 the ISSUE FEE shown above.

II. PART B - FEE(S) TRANSMITTAL, or its equivalent, must be completed and returned to the United States Patent and Trademark Office (USPTO) with your ISSUE FEE and PUBLICATION FEE (if required). If you are charging the fee(s) to your deposit account, section "4b" of Part B - Fee(s) Transmittal should be completed and an extra copy of the form should be submitted. If an equivalent of Part B is filed, a request to reapply a previously paid issue fee must be clearly made, and delays in processing may occur due to the difficulty in recognizing the paper as an equivalent of Part B.

III. All communications regarding this application must give the application number. Please direct all communications prior to issuance to Mail Stop ISSUE FEE unless advised to the contrary.

IMPORTANT REMINDER: Utility patents issuing on applications filed on or after Dec. 12, 1980 may require payment of maintenance fees. It is patentee's responsibility to ensure timely payment of maintenance fees when due.

#### PART B - FEE(S) TRANSMITTAL

### Complete and send this form, together with applicable fee(s), to: Mail Mail Stop ISSUE FEE

Commissioner for Patents P.O. Box 1450 Alexandria, Virginia 22313-1450

or <u>Fax</u> (571)-273-2885

INSTRUCTIONS: This form should be used for transmitting the ISSUE FEE and PUBLICATION FEE (if required). Blocks 1 through 5 should be completed where ar in m

ppropriate. All further adicated unless correcteraintenance fee notificate	correspondence includin ed below or directed oth	g the Patent, advance of the lerwise in Block 1, by (a	rders and notification of a) specifying a new co	of ma	aintenance fees wondence address;	ill be and/or	mailed to the current (b) indicating a sepa	correspo rate "FE	ndence address as E ADDRESS" for
	ENCE ADDRESS (Note: Use Blo	ock 1 for any change of address)	Ŧ	Fee(s	) Transmittal This	s certif	can only be used for icate cannot be used for such as an assignment ling or transmission.	or any of	her accompanying
400 GARDEN C SUITE 300					Cert	ificate	of Mailing or Transi s) Transmittal is being ficient postage for firs ISSUE FEE address 1) 273-2885, on the da	nission	
GARDEN CITY	, NY 11530								(Depositor's name)
									(Signature)
			L						(Date)
APPLICATION NO.	LICATION NO. FILING DATE FIRST NAMED INVE			NTOR ATTORNEY DOCKET NO. CONFIRMATION NO.				RMATION NO.	
10/705,614 ITLE OF INVENTION	11/10/2003 : METHODS OF ENHA	NCING RADIATION E	James F. Hainfeld FFECTS WITH META	LΝ	ANOPARTICLES		16049Z		4692
APPLN. TYPE	SMALL ENTITY	ISSUE FEE DUE	PUBLICATION FEE DU	UE	PREV. PAID ISSUE	FEE	TOTAL FEE(S) DUE		DATE DUE
nonprovisional	YES	\$755	\$300		\$0		\$1055		03/31/2009
EXAM	EXAMINER ART UNIT CLASS-SUBCLA		CLASS-SUBCLASS	s					
HOPKINS, CI	HRISTINE D	3735	600-001000	_					
Change of correspondence address or indication of "Fee Address" (37 FR 1.363).  Change of correspondence address (or Change of Correspondence Address form PTO/SB/122) attached.  "Fee Address" indication (or "Fee Address" Indication form PTO/SB/47; Rev 03-02 or more recent) attached. Use of a Customer Number is required.			(1) the names of up or agents OR, alterr (2) the name of a si registered attorney 2 registered patent a	2. For printing on the patent front page, list  (1) the names of up to 3 registered patent attorneys or agents OR, alternatively,  (2) the name of a single firm (having as a member a registered attorney or agent) and the names of up to 2 registered patent attorneys or agents. If no name is listed, no name will be printed.					
PLEASE NOTE: Unl recordation as set forth (A) NAME OF ASSIG	less an assignee is identi h in 37 CFR 3.11. Comp GNEE	A TO BE PRINTED ON The field below, no assignee oletion of this form is NO categories (will not be presented to the presented of the presented	data will appear on th T a substitute for filing (B) RESIDENCE: (Cl	e pat an a	ent. If an assignessignment. and STATE OR C	OUNT			
a. The following fee(s) a  Issue Fee  Publication Fee (N		4lpermitted)	b. Payment of Fee(s): (I A check is enclose Payment by credit The Director is her	Pleas ed. card	e first reapply an  Form PTO-2038 authorized to charge	y prev	iously paid issue fee s	hown al	bove) or credit any
a. Applicant claims	<b>tus</b> (from status indicated s SMALL ENTITY statu	s. See 37 CFR 1.27.		_	-		FITY status. See 37 CF	-	
OTE: The Issue Fee and terest as shown by the r	d Publication Fee (if requeecords of the United Sta	iired) will not be accepte tes Patent and Trademark	d from anyone other the Office.	an th	e applicant; a regis	stered a	nttorney or agent; or th	e assigne	e or other party in
Authorized Signature					Date				
Typed or printed name	e				Registration N	o			
his collection of inform n application. Confident Ibmitting the completed its form and/or suggesti ox 1450, Alexandria, V Ilexandria, Virginia 223	tiality is governed by 35 dapplication form to the tons for reducing this bur riginia 22313-1450. DO	FR 1.311. The informatic U.S.C. 122 and 37 CFR USPTO. Time will vary den, should be sent to th O NOT SEND FEES OR	on is required to obtain 1.14. This collection is depending upon the ine Chief Information Of COMPLETED FORMS	or re estindivid fficer TO	tain a benefit by th nated to take 12 n dual case. Any co , U.S. Patent and THIS ADDRESS	ne publ ninutes mment Fraden . SENI	ic which is to file (and to complete, includin s on the amount of tin ark Office, U.S. Depa D TO: Commissioner f	by the Ug gatherine you reartment of Paten	JSPTO to process) ng, preparing, and equire to complete of Commerce, P.O. ts, P.O. Box 1450,

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it displays a valid OMB control number.



# United States Patent and Trademark Office

UNITED STATES DEPARTMENT OF COMMERCE United States Patent and Trademark Office Address: COMMISSIONER FOR PATENTS P.O. Box 1450 Alexandria, Virginia 22313-1450 www.uspto.gov

APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.	
10/705,614	11/10/2003	James F. Hainfeld	16049Z	4692	
23389 75	590 12/31/2008		EXAM	INER	
SCULLY SCOT	Γ MURPHY & PRES	HOPKINS, CHRISTINE D			
400 GARDEN CIT	TY PLAZA	ART UNIT	PAPER NUMBER		
SUITE 300 GARDEN CITY, NY 11530			3735 DATE MAILED: 12/31/2008		

## **Determination of Patent Term Adjustment under 35 U.S.C. 154 (b)**

(application filed on or after May 29, 2000)

The Patent Term Adjustment to date is 574 day(s). If the issue fee is paid on the date that is three months after the mailing date of this notice and the patent issues on the Tuesday before the date that is 28 weeks (six and a half months) after the mailing date of this notice, the Patent Term Adjustment will be 574 day(s).

If a Continued Prosecution Application (CPA) was filed in the above-identified application, the filing date that determines Patent Term Adjustment is the filing date of the most recent CPA.

Applicant will be able to obtain more detailed information by accessing the Patent Application Information Retrieval (PAIR) WEB site (http://pair.uspto.gov).

Any questions regarding the Patent Term Extension or Adjustment determination should be directed to the Office of Patent Legal Administration at (571)-272-7702. Questions relating to issue and publication fee payments should be directed to the Customer Service Center of the Office of Patent Publication at 1-(888)-786-0101 (571)-272-4200.

	Application No.	Applicant(s)	
	10/705,614	HAINFELD ET AL.	
Notice of Allowability	Examiner	Art Unit	
	CHRISTINE D. HOPKINS	3735	
	CHRISTINE D. HOFKINS	3733	
The MAILING DATE of this communication appeal All claims being allowable, PROSECUTION ON THE MERITS IS herewith (or previously mailed), a Notice of Allowance (PTOL-85) NOTICE OF ALLOWABILITY IS NOT A GRANT OF PATENT RIOF of the Office or upon petition by the applicant. See 37 CFR 1.313	(OR REMAINS) CLOSED in this or other appropriate communical GHTS. This application is subject	application. If not included ion will be mailed in due course. <b>THIS</b>	
1. $\boxtimes$ This communication is responsive to <u>the Amendment filed</u>	<u>18 December 2008</u> .		
2. The allowed claim(s) is/are <u>1-22,25,69 and 71-89</u> .			
3. $\square$ Acknowledgment is made of a claim for foreign priority ur	nder 35 U.S.C. § 119(a)-(d) or (f).		
a) All b) Some* c) None of the:			
<ol> <li>Certified copies of the priority documents have</li> </ol>	e been received.		
<ol><li>Certified copies of the priority documents have</li></ol>	• •		
<ol><li>Copies of the certified copies of the priority do</li></ol>	cuments have been received in the	nis national stage application from the	
International Bureau (PCT Rule 17.2(a)).			
* Certified copies not received:			
Applicant has THREE MONTHS FROM THE "MAILING DATE" noted below. Failure to timely comply will result in ABANDONN THIS THREE-MONTH PERIOD IS NOT EXTENDABLE.		oly complying with the requirements	
4. A SUBSTITUTE OATH OR DECLARATION must be subm INFORMAL PATENT APPLICATION (PTO-152) which give			
5. CORRECTED DRAWINGS ( as "replacement sheets") mus	st be submitted.		
(a) ☐ including changes required by the Notice of Draftspers	on's Patent Drawing Review ( P1	O-948) attached	
1) 🔲 hereto or 2) 🔲 to Paper No./Mail Date	•		
(b) ☐ including changes required by the attached Examiner's Paper No./Mail Date	s Amendment / Comment or in th	e Office action of	
Identifying indicia such as the application number (see 37 CFR 1 each sheet. Replacement sheet(s) should be labeled as such in t			
6. DEPOSIT OF and/or INFORMATION about the depo attached Examiner's comment regarding REQUIREMENT			
Attachment(s) 1. ☐ Notice of References Cited (PTO-892)	5. ☐ Notice of Informa	al Patent Application	
2.  Notice of Draftperson's Patent Drawing Review (PTO-948)	6. Interview Summa		
3. Information Disclosure Statements (PTO/SB/08),	Paper No./Mail 7.	Date ndment/Comment	
Paper No./Mail Date  4.  Examiner's Comment Regarding Requirement for Deposit  8.  Examiner's Statement of Reasons for Allowance of Biological Material			
-	9.  Other		

## **DETAILED ACTION**

1. This Office Action is responsive to the Amendment filed 18 December 2008. Claims 1-22, 25, 69 and 71-89 are now pending.

## Allowable Subject Matter

2. Claims 1-22, 25, 69, and 71-89 are allowable over the prior art of record. The following is an examiner's statement of reasons for allowance: regarding claims 1-22 and 25, the prior art of record does not teach or fairly suggest a method of enhancing the effects of radiation in an animal or a method of ablating a tissue or a population of cells in an animal as disclosed by Applicant in claims 1 and 2, wherein the metal nanoparticles are administered in an amount to achieve a concentration in the animal of at least about 0.1% metal by weight. While the prior art teaches a concentration of the metal elements in the nanoparticles, the prior art does not disclose a percentage of metal *in the target tissue*.

Regarding claims 69 and 71-87, the prior art of record does not teach or fairly suggest a method of enhancing the effects of radiation in an animal as disclosed by Applicant, wherein the metal nanoparticles are administered in an amount to achieve a concentration of *metal* in the tissue of the animal of at least about 0.1% by weight.

Regarding claim 88, the prior art of record does not teach or fairly suggest a method of enhancing the effects of radiation in an animal as disclosed by Applicant, comprising administering an amount of metal nanoparticles to said animal and subsequently irradiating the animal with radiation wherein said nanoparticles comprise a

surface layer material comprising thioglucose. Regarding claim 89, the prior art of record does not teach or fairly suggest a method of enhancing the effects of radiation in an animal as disclosed by Applicant, comprising administering an amount of metal nanoparticles to said animal and subsequently irradiating the animal with radiation wherein said nanoparticles are polyanions of metals complexed with quaternary ammonium salts.

#### Terminal Disclaimer

3. The terminal disclaimer filed on 18 December 2008 disclaiming the terminal portion of any patent granted on this application which would extend beyond the expiration date of either U.S. Patent No. 6,955,639 or U.S. Patent No. 7,367,934 has been reviewed and is accepted. The terminal disclaimer has been recorded.

### Conclusion

Any comments considered necessary by applicant must be submitted no later than the payment of the issue fee and, to avoid processing delays, should preferably accompany the issue fee. Such submissions should be clearly labeled "Comments on Statement of Reasons for Allowance."

Any inquiry concerning this communication or earlier communications from the examiner should be directed to CHRISTINE D. HOPKINS whose telephone number is (571)272-9058. The examiner can normally be reached on Monday-Friday, 7 a.m.-3:30 p.m..

Application/Control Number: 10/705,614 Page 4

Art Unit: 3735

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Charles Marmor, II can be reached on (571) 272-4730. The fax phone number for the organization where this application or proceeding is assigned is 571-273-8300.

Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see http://pair-direct.uspto.gov. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free). If you would like assistance from a USPTO Customer Service Representative or access to the automated information system, call 800-786-9199 (IN USA OR CANADA) or 571-272-1000.

/Charles A. Marmor, II/ Supervisory Patent Examiner Art Unit 3735

/C. D. H./ Christine D Hopkins Examiner Art Unit 3735